# **PCT**

## INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference J1557-PCT	FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.				
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)			
PCT/EP 00/06245	03/07/2000	03/07/1999			
JANSSEN PHARMACEUTICA N.V	1.				
according to Article 18. A copy is being to This International Search Report consists					
Basis of the report					
a. With regard to the language, the	international search was carried out on the ba less otherwise indicated under this item.	sis of the international application in the			
the international search v Authority (Rule 23.1(b)).	was carried out on the basis of a translation of t	he international application furnished to this			
was carried out on the basis of the contained in the internation of the filed together with the internation of the furnished subsequently to the statement that the subsequent that the su					
the statement that the inf furnished	ormation recorded in computer readable form i	s identical to the written sequence listing has been			
Certain claims were for     Unity of invention is lace.	und unsearchable (See Box I).				
onity of invention is tac	oring (see box II).	•			
4. With regard to the <b>title</b> ,					
	ubmitted by the applicant.	•			
the text has been established.	shed by this Authority to read as follows:				
the text has been establis	ubmitted by the applicant. shed, according to Rule 38.2(b), by this Authori e date of mailing of this international search rep				
6. The figure of the <b>drawings</b> to be pub	lished with the abstract is Figure No.	1			
X as suggested by the appl		None of the figures.			
because the applicant fai	•				
Decause this rigure better	r characterizes the invention.				

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A. CLASSIFICATION OF SUBJECT MATTER IPC 7 G01N35/04 G02B21/00

According to International Patent Classification (IPC) or to both national classification and IPC

#### B. FIELDS SEARCHED

 $\begin{array}{ll} \mbox{Minimum documentation searched (classification system followed by classification symbols)} \\ \mbox{IPC 7} & \mbox{B65D} & \mbox{G02B} & \mbox{G01N} \end{array}$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

#### EPO-Internal

C. DOCUMI	C. DOCUMENTS CONSIDERED TO BE RELEVANT							
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.						
Α	WO 98 52047 A (AURORA BIOSCIENCES CORP) 19 November 1998 (1998-11-19) page 84, paragraph 3 -page 85, paragraph 2; figure 15	1,2						
Α	WO 98 59033 A (SANKYO CO ;ICHIKAWA MASATO (JP); INABA KAZUHIRO (JP); NITTETSU MIN) 30 December 1998 (1998-12-30) figure 6	2						
A	DE 298 06 303 U (ARCHYTAS AUTOMATION GMBH) 3 September 1998 (1998-09-03) page 11, last paragraph -page 12, paragraph 2; figure 1 page 16, line 10 - line 32; figure 4	1,2,6,10						
Î								

X Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
<ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the art which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the international filing date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filing date but later than the priority date claimed</li> </ul>	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  "&" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
25 October 2000	02/11/2000
Name and mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer  Hocquet, A

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Int mal Application No PCT/EP 00/06245

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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevan	t to claim No.
Α	EP 0 245 920 A (CAMBRIDGE LIFE SCIENCES) 19 November 1987 (1987-11-19) column 4, line 40 -column 6, line 13; figures 2,3	6	,7
Α	US 5 386 318 A (KUEHNERT JUERGEN ET AL) 31 January 1995 (1995-01-31) cited in the application column 3, line 48 -column 4, line 42; figures		,2,7,8, 0
A	WO 97 39348 A (ALPHA SCIENT CORP) 23 October 1997 (1997-10-23) cited in the application page 7, last paragraph -page 8, paragraph 3; figures 3-6	1	,2,10
A	DE 297 12 535 U (BODENSEEWERK PERKIN ELMER CO) 18 September 1997 (1997-09-18) cited in the application page 6, paragraph 2; figure 1 page 7, paragraph 3; figures 2,4	1	,2,10
A	WO 93 12430 A (BAXTER DIAGNOSTICS INC) 24 June 1993 (1993-06-24) page 37, line 23 -page 39, line 33; figures 26-31	1	,2,10
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### INTERMINIONAL SEARCH REPORT

form on patent family members

Internal Application No PCT/EP 00/06245

	itent document I in search report		Publication date	Patent family Publication member(s) date
WO	9852047	Α	19-11-1998	US 5985214 A 16-11-1999 AU 7478398 A 08-12-1998 EP 1010009 A 21-06-2000
WO	9859033	A	30-12-1998	JP 11009260 A 19-01-1999 AU 8037798 A 04-01-1999 EP 1018544 A 12-07-2000
DE	29806303	U	03-09-1998	NONE
EP	0245920	A	19-11-1987	GB 2190195 A 11-11-1987 AU 6985487 A 12-11-1987 DK 236487 A 10-11-1987 FI 872019 A 10-11-1987 HU 43733 A,B 30-11-1987 JP 63061142 A 17-03-1988 US 4810096 A 07-03-1989
US	5386318	Α	31-01-1995	DE 4131360 A 25-03-1993 WO 9306516 A 01-04-1993
WO	9739348	Α	23-10-1997	AU 2600897 A 07-11-1997
DE	29712535	U	18-09-1997	NONE
WO	9312430	Α	24-06-1993	AU 3333493 A 19-07-1993 CA 2101951 A 19-07-1992 EP 0572640 A 08-12-1993 JP 6507976 T 08-09-1994

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### INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)



Applicant's or agent's file reference J1557-PCT			FOR FURTHER ACTION		cation of Transmittal of Interr y Examination Report (Form	
Internation	al app	lication No.	International filing date (day/month	/year)	Priority date (day/month/)	year)
PCT/EP			03/07/2000	•	03/07/1999	
Internation G01N35		ent Classification (IPC) or na	tional classification and IPC		1	
Applicant JANSSE	N PH	HARMACEUTICA N.V.				
1. This and i	intern s tran	ational preliminary exami smitted to the applicant a	ination report has been prepared according to Article 36.	l by this Inte	ernational Preliminary Ex	camining Authority
2. This	REPO	ORT consists of a total of	6 sheets, including this cover s	heet.		·
i	oeen a	amended and are the bas	d by ANNEXES, i.e. sheets of th sis for this report and/or sheets of O7 of the Administrative Instructi	ontaining re	ectifications made before	js which have this Authority
There		ist of a total of	2 chests			•
ines	e ann	exes consist of a total of	Z Stieets.			
1	Ø	Basis of the report	ating to the following items:			
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III IV			pinion with regard to novelty, inv	enuve step	and industrial applicabili	щ
V		Reasoned statement ur	nder Article 35(2) with regard to ons suporting such statement	novelty, inv	entive step or industrial a	applicability;
VI		Certain documents cite	ed			
VII	$\boxtimes$	Certain defects in the ir	nternational application			
VIII	Ø	Certain observations or	n the international application			
Date of su	bmissi	on of the demand	Date of	completion of	f this report	
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# INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/EP00/06245

I.	Bas	is fth report						
1.	With regard to the <b>elements</b> of the international application (Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17)): <b>Description, pages:</b>							
	1-1	1	as originally filed					
	Cla	ims, No.:						
	1-15	5	as received on	26/06/2001	with letter of	26/06/2001		
	Dra	wings, sheets:						
	1/3-	3/3	as originally filed					
2.	With lang	n regard to the <b>lang</b> guage in which the	guage, all the elements marked international application was file	above were a	vailable or furnished t erwise indicated under	o this Authority in the r this item.		
	The	se elements were	available or furnished to this Au	thority in the fo	ollowing language: ,	which is:		
		the language of a	translation furnished for the pur	poses of the i	nternational search (u	nder Rule 23.1(b)).		
			ublication of the international ap					
		the language of a 55.2 and/or 55.3).	translation furnished for the pur	poses of inter	national preliminary e	xamination (under Rule		
3.	With inte	n regard to any <b>nuo</b> rnational prelimina	cleotide and/or amino acid sec ry examination was carried out o	<b>quence</b> disclo on the basis o	sed in the internationa f the sequence listing:	al application, the		
		contained in the ir	nternational application in writter	n form.				
		filed together with	the international application in	computer read	lable form.			
		furnished subsequ	uently to this Authority in written	form.		•		
		furnished subsequ	uently to this Authority in compu	ter readable fo	orm.			
			at the subsequently furnished wi application as filed has been furr		e listing does not go b	eyond the disclosure in		
		The statement that listing has been fu	at the information recorded in co urnished.	mputer reada	ble form is identical to	the written sequence		

□ the description,
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☐ the claims,

4. The amendments have resulted in the cancellation of:

pages:

Nos.:

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# INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/EP00/06245

		the drawings,	sheets:					
5.		☐ This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):						
		(Any replacement sh report.)	eet contair	ning such	amendments must be referred to under item 1 and annexed to this			
6.	Add	itional observations, if	f necessar	y:				
V.		soned statement un tions and explanatio			ith regard to novelty, inventive step or industrial applicability; h statement			
1.	Stat	ement						
	Nov	elty (N)	Yes: No:	Claims Claims	1-15			
	Inve	entive step (IS)	Yes:	Claims	1-15			

2. Citations and explanations see separate sheet

Industrial applicability (IA)

#### VII. Certain defects in the international application

The following defects in the form or contents of the international application have been noted: see separate sheet

Claims

Claims

Claims 1-15

No:

Yes:

No:

#### VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made: see separate sheet

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#### Re Item V

Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

- The following documents are referred to in this report: 1.
  - D1: WO 98 52047 A (AURORA BIOSCIENCES CORP) 19 November 1998 (1998-11-19)
  - D2: WO 98 59033 A (SANKYO CO ;ICHIKAWA MASATO (JP); INABA KAZUHIRO (JP); NITTETSU MIN) 30 December 1998 (1998-12-30)
  - D3: DE 298 06 303 U (ARCHYTAS AUTOMATION GMBH) 3 September 1998 (1998-09-03)
  - D4: EP-A-0 245 920 (CAMBRIDGE LIFE SCIENCES) 19 November 1987 (1987-11-19)
  - D5: US-A-5 386 318 (KUEHNERT JUERGEN ET AL) 31 January 1995 (1995-01-31) cited in the application
  - D6: WO 97 39348 A (ALPHA SCIENT CORP) 23 October 1997 (1997-10-23) cited in the application
  - D7: DE 297 12 535 U (BODENSEEWERK PERKIN ELMER CO) 18 September 1997 (1997-09-18) cited in the application
  - D8: WO 93 12430 A (BAXTER DIAGNOSTICS INC) 24 June 1993 (1993-06-24)

#### 2. Review of the prior art documents:

D1 describes a complex automated system (see e.g. fig 5) for well plates for use in chemical analysis, having an area for storage and retrieval (302, 303 at rear of fig. 5), and transporting means for removing well plates and transporting them to analysis/processing stations. Page 31, lines 5-7, states that plates are taken from a stack and presented to a transporter in an exact order. This seems to be the idea of singulation. Page 84, lines 22-25, discloses the idea of removing plates from the bottom of a stack in a vertical direction., Fig. 15B and page 85, second paragraph describes how plates are removed downwards vertically. The lower part of fig. 15B seems to show a platform by which plates are lowered further, and can be considered as a "singulator".

D2 relates to a stacker for laboratory dishes. As shown in fig. 6, particularly 6c, it would

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appear that the plates fall downwardly and are removed sideways. There appears to be no device acting as a "singulator" which stops at an "intermediate" position.

D3 describes a system in which the well plates are stored in racks on columns of trays. The trays can be moved vertically to bring one in line with the testing/processing apparatus, and the tray is pulled out to position it next to the testing/processing head.

D4 describes a system in which a well plate is mounted in a drawer which can be slid into the testing apparatus housing.

D5 discloses a system in which specimens 6 are stacked in a cassette (fig. 5a), a specimen being removed for inspection by pushing it using a slide 12 onto a rotary arm 5.

D6 discloses an arrangement of belts having treads supporting a column of slides therebetween (figs. 10-13). As a slide reaches the bottom of the belt structure, it falls out of the treads onto a shelf, which is rotated to invert the slide.

D7 describes a magazine for sample holders, the magazine being moved vertically until a position is reached for removal of a desired holder, by sideways movement.

D8 discloses a transporting mechanism for carriers (see figs 26-31, and pages 38-40), in which a platform 70, which could be considered to be a "singulator", receives the bottom carrier of a pile, which is supported by pins 150. The platform lifts the bottom carrier and the rest of the pile, and the pins are released. The platform then lowers the pile, until pins 150 move in again to support the pile above the bottom one. The removed carrier is then moved on the platform to a hub access 23 and then to a hub access 25 thereunder.

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#### 3. Novelty and inv ntiv st p:

<u>The invention relates to</u> a transporter for transporting specimen carriers to a processing station and a method of transporting.

Object of the invention is to provide a compact system with analysis facility.

Solution provided by the invention is as defined in claims 1 and 10: input and output magazines have a singulator therebetween bringing carriers one by one to an intermediate position between the magazines.

#### Prior art documents:

Claim 1 differs from D1 in that a second magazine is provided, under the first, and that a singulator halt position in between the two magazines is specified.

Claim 1 differs from D8 in that a second magazine is provided, under the first, and that a singulator halt position is between the two magazines.

The other documents seem to be less relevant than D1 and D8.

Thus claim 1 is novel in the light of the prior art. There is no hint in the prior art towards these new features, so that the subject matter of claim 1 is inventive.

Claim 10 is of roughly equivalent scope to claim 1, and so is also novel and inventive. The dependent claims are likewise novel and inventive.

#### Re Item VII

#### Certain defects in the international application

Prior art documents D1, D6 and D8, at least, should have been referred to.

#### Re Item\_VIII

#### Certain observations on the international application

- 1. Although the statement of invention on page 3, lines 13-18 corresponds in scope with claim 10, that on page 2, line 2, onwards, does not properly correspond with the wording of claim 1, leading to lack of consistency and clarity.
- 2. The references to an aspect of the invention on page 6, lines 8-9, and the spirit of the invention on page 11, line 8, are confusing.

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#### **CLAIMS**

1. A transporter for transporting carriers to a processing station, comprising:
a first vertical input magazine of carriers, a second output magazine of carriers, the first
and second magazines being located one above the other in either order; and
a singulator located between the output of the first magazine and the input of the second
magazine, the singulator being adapted to receive one carrier at a time from the first
magazine and to move the carrier from the first magazine vertically to a position
intermediate the first and the second magazines and to halt the carrier at this position.

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- 2. The transporter according to claim 1, wherein the singulator is adapted to move a received carrier incrementally upwards or downwards at the intermediate position independent of any movement of the carriers in the first and/or second magazines.
- 3. The transporter according to claim 2, wherein the incremental movement comprises autofocussing about a processing position.
- The transporter according to any previous claim wherein the singulator comprises a
  release mechanism and a conveyor, the release mechanism releasing carriers from the first
  magazine one-at-a-time to the conveyor.
  - 5. The transporter according to any of the previous claims wherein the conveyor of the singulator comprises a pair of opposed belts, the belts having external protuberances or ledges and the belts being arranged vertically, so that the external ledges from the two belts define horizontally-oriented shelves which are capable of holding a received carrier in a substantially horizontal orientation.
  - 6. The transporter according to any previous claim, further comprising a processing tool movable horizontally to a location above and/or below the carrier when the carrier is at the intermediate position in the singulator.
  - 7. The transporter according to claim 6, wherein the processing tool is an optical receiver for capturing an image, in particular a video image of the contents of the carrier.

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#### Bird Goën & Co

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26 June 2001

- 8. The transporter according to claim 7, wherein the optical receiver comprises a microscope.
- 9. The transporter according to any previous claim, wherein the carriers are Petri dishes, multi-well plates or any other device containing biological specimens.
  - 10. A method of transporting specimen carriers to a processing position, comprising the steps of:
- providing a vertical input magazine and an output magazine of carriers, one above the other;

singling out a carrier from the input magazine and moving it vertically to an intermediate position which is located between the input and output magazines and halting it there; and moving the carrier to an output location where it is transferred to the output magazine.

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- 11. The method according to claim 10, further comprising moving the carrier incrementally up or down at the intermediate position using the singulator.
- 12. The method according to claim 11, the step of moving the carrier incrementally comprising autofocussing about a processing position.
  - 13. The method according to any one of claims 10 to 12, further comprising the steps of: moving a processing tool horizontally to a position above and/or below the carrier at the intermediate position; and
- 25 withdrawing the processing tool after carrying out processing.
  - 14. The method according to any of claims 10 to 13, wherein the movement of the carrier to the intermediate position is independent of movements of the carriers in the input and output magazines

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15. The method according to claim 13 or 14, wherein the processing tool is an optical receiver.

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#### P. ENT COOPERATION TREAT

### From the INTERNATIONAL BUREAU **PCT** Commissioner **NOTIFICATION OF ELECTION US Department of Commerce** United States Patent and Trademark (PCT Rule 61.2) Office, PCT 2011 South Clark Place Room CP2/5C24 Arlington, VA 22202 **ETATS-UNIS D'AMERIQUE** Date of mailing (day/month/year) in its capacity as elected Office 15 February 2001 (15.02.01) International application No. Applicant's or agent's file reference PCT/EP00/06245 s1557-PCT International filing date (day/month/year) Priority date (day/month/year) 03 July 2000 (03.07.00) 03 July 1999 (03.07.99) **Applicant** NUYENS, Roger, François, Gabrielle, Armand 1. The designated Office is hereby notified of its election made: X in the demand filed with the International Preliminary Examining Authority on: 10 January 2001 (10.01.01) in a notice effecting later election filed with the International Bureau on: 2. The election made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b). Authorized officer The International Bureau of WIPO 34, chemin des Colombettes A. Karkachi 1211 Geneva 20, Switzerland

Telephone No.: (41-22) 338.83.38

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## PATENT COOPERATION TREATY

# PCT

### INTERNATIONAL SEARCH REPORT

	(PCT Article 18 and Rule	s 43 and 44)			
Applicant's or agent's file refere	i ionioninen '	ee Notification of Tra	insmittal of International Search Report		
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International application No.	International filing date (day)	month/year) (E	arliest) Priority Date (day/month/year)		
PCT/EP 00/06245	03/07/200		03/07/1999		
Applicant	03/07/200	<u> </u>	03/07/1999		
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JANSSEN PHARMACEUT	ICA N.V.				
This International Search Repo	ort has been prepared by this International	Searching Authority	and in transmitted to the section		
according to Article 18. A copy	is being transmitted to the International B	ureau.	and is transmitted to the applicant		
This International Secret Sec	2		•		
This International Search Repo	ort consists of a total of3 npanied by a copy of each prior art docum	_ sheets.			
		ent cited in this repon	τ.		
Basis of the report					
a. With regard to the land language in which it was	guage, the international search was carried as filed, unless otherwise indicated under t	d out on the basis of t	the international application in the		
Authority (Rule	al search was carried out on the basis of a $23.1(b)$ .	translation of the inte	ernational application furnished to this		
b. With regard to any nuc	cleotide and/or amino acid sequence dis	closed in the internati	ional application, the international search		
	basis of the sequence listing: e international application in written form.				
	rith the international application in compute	er readable form.			
	equently to this Authority in written form.				
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the statement to	hat the subsequently furnished written seq plication as filed has been furnished.	uence listing does no	t go beyond the disclosure in the		
		eadable form is identi	cal to the written sequence listing has been		
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2. Certain claims	were found unsearchable (See Box I).				
	tion is lacking (see Box II).				
4. With regard to the title,					
	oved as submitted by the applicant.				
the text has bee	en established by this Authority to read as	follows:			
5. With regard to the abstract,			·		
X the text is appro	ved as submitted by th applicant.				
th text has bee	n established, according to Rule 38.2(b)	by this Authority as it a	appears in Box III. The applicant may,		
	h from the date of mailing of this internation to be published with the abstract is Figure	nai search report, sub	ornit comments to this Authority.		
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	plicant failed to suggest a figure.		Non of the figures.		
	are better characterizes the invention.				

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#### INTERN NAL SEARCH REPORT



CLASSIFICATION OF SUBJECT MATTER PC 7 G01N35/04 G02B G02B21/00 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) B65D G02B G01N IPC 7 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with Indication, where appropriate, of the relevant passages Category \* 1,2 WO 98 52047 A (AURORA BIOSCIENCES CORP) Α 19 November 1998 (1998-11-19) page 84, paragraph 3 -page 85, paragraph 2; figure 15 2 WO 98 59033 A (SANKYO CO ; ICHIKAWA MASATO Α (JP); INABA KAZUHIRO (JP); NITTETSU MIN) 30 December 1998 (1998-12-30) figure 6 1,2,6,10 DE 298 06 303 U (ARCHYTAS AUTOMATION GMBH) Α 3 September 1998 (1998-09-03) page 11, last paragraph -page 12, paragraph 2; figure 1 page 16, line 10 - line 32; figure 4 Patent family members are listed in annex. Further documents are listed in the continuation of box C. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance invention "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention filing date cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such document. "O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled other means in the art. "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of mailing of the international search report Date of the actual completion of the international search 02/11/2000 25 October 2000 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Hocquet, A Fax: (+31-70) 340-3016

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	Jation) DOCUMENTS CONSIDERED TO BE RELEVANT	Dalaman at the At
ategory *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
4	EP 0 245 920 A (CAMBRIDGE LIFE SCIENCES) 19 November 1987 (1987-11-19) column 4, line 40 -column 6, line 13; figures 2,3	6,7
A	US 5 386 318 A (KUEHNERT JUERGEN ET AL) 31 January 1995 (1995-01-31) cited in the application column 3, line 48 -column 4, line 42; figures	1,2,7,8,
A	WO 97 39348 A (ALPHA SCIENT CORP) 23 October 1997 (1997-10-23) cited in the application page 7, last paragraph -page 8, paragraph 3; figures 3-6	1,2,10
Α.	DE 297 12 535 U (BODENSEEWERK PERKIN ELMER CO) 18 September 1997 (1997-09-18) cited in the application page 6, paragraph 2; figure 1 page 7, paragraph 3; figures 2,4	1,2,10
А	WO 93 12430 A (BAXTER DIAGNOSTICS INC) 24 June 1993 (1993-06-24) page 37, line 23 -page 39, line 33; figures 26-31	1,2,10
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	atent document d in search repor	t	Publication dat	Patent family Publication member(s) date
WO	9852047	Α	19-11-1998	US 5985214 A 16-11-1999 AU 7478398 A 08-12-1998 EP 1010009 A 21-06-2000
WO	9859033	A	30-12-1998	JP 11009260 A 19-01-1999 AU 8037798 A 04-01-1999 EP 1018544 A 12-07-2000
DE	29806303	U	03-09-1998	NONE
EP	0245920	A	19-11-1987	GB 2190195 A 11-11-1987 AU 6985487 A 12-11-1987 DK 236487 A 10-11-1987 FI 872019 A 10-11-1987 HU 43733 A,B 30-11-1987 JP 63061142 A 17-03-1988 US 4810096 A 07-03-1989
US	5386318	Α	31-01-1995	DE 4131360 A 25-03-1993 WO 9306516 A 01-04-1993
WO	9739348	Α	23-10-1997	AU 2600897 A 07-11-1997
DE	29712535	U	18-09-1997	NONE
WO	9312430	A	24-06-1993	AU 3333493 A 19-07-1993 CA 2101951 A 19-07-1992 EP 0572640 A 08-12-1993 JP 6507976 T 08-09-1994

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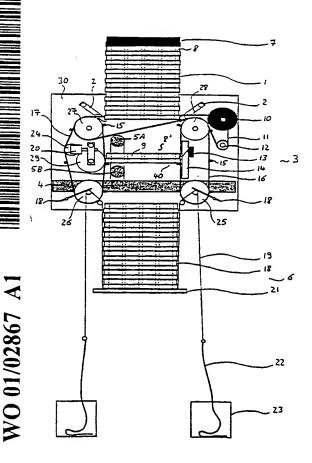
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(54) Title: APPARATUS FOR TRANSPORTING CARRIERS TO A PROCESSING STATION AND METHOD OF OPERATING THE SAME



(57) Abstract: The present application a transporter (100) for transporting specimen carriers (8) to a processing station (40), comprising a first vertical input magazine (1) of carriers (8), a second vertical output magazine (6) of carriers, and a singulator (3) located between the first and second magazines (1, 6). The first and second magazines (1, 6) may be located one above the other. Preferably, the first and second magazines (1, 6) and the singulator (3) are vertically aligned so that a carrier (8) follows a straight vertical path. This provides a compact arrangement which takes up the minimum of floor space in the laboratory. The singulator (3) receives carriers (8) one-at-a-time from the input magazine (1) and moves them vertically to a processing position (40). After processing, the singulator (3) then delivers each carrier to an output location where it is transferred to the output magazine (6). The singulator (3) may move a carrier (8) upwards or downwards at the processing position (40) independent of any movement of the carriers (8) in the input and/or output magazines (1, 6). The drives for the input and output magazine (1, 6) are preferably simply gravity drives but the present invention is not limited thereto. The drive (10) for the singulator is preferably a stepping motor or DC servo-motor.



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## APPARATUS FOR TRANSPORTING CARRIERS TO A PROCESSING STATION AND METHOD OF OPERATING THE SAME

The present invention relates to handling of objects, particularly objects to be scanned optically or viewed as for instance microscope slides, specimen carriers, trays or Petri dishes. In particular, the present invention relates to an apparatus and method for transporting carriers, such as microscope slides or trays to a processing position..

## TECHNICAL BACKGROUND OF THE INVENTION

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Devices for conveying specimens particularly for clinical laboratories are known from DE-U-297 12 535, WO 97/39348, WO 93/06516. However, most of these devices need to grasp the specimen, or push the specimen sideways out of a vertical magazine to a viewing position or need to move them in some other way to a viewing or processing position. It is often inconvenient grasp or push a delicate specimen holder such as a microscope slide, either because the slide is fragile, or because the specimen, or the indicia imprinted on the slide, can easily be abraded and thus obliterated. In addition, the known devices require a separate mechanism for autofocusing onto the specimen to be viewed. For example, autofocusing mechanisms are known from WO 96/01438, US 3,721,759, WO 96/10196, US 5,790,692, US 5,790,710. These require a stage for moving the specimen in at least two dimensions and often in three. This stage is in addition to the vertical delivery systems described above. Each specimen must be conveyed to a first position with one mechanism, transferred to a viewing position with another mechanism and then autofocused using yet another mechanism. This makes the combination expensive, bulky and complicated.

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It is an object of the present invention to provide a compact system and method for delivery of specimens, particularly those to be viewed optically, to a processing or viewing position.

It is a further object of the present invention to provide an apparatus and method for obtaining an autofocus position for an optical instrument in a short time.

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It is a further object of the present invention to provide an apparatus and method for obtaining an autofocus position for an optical instrument which is simple, compact and less expensive than known apparatus and methods.

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### SUMMARY OF THE INVENTION

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One aspect of the present invention is to integrate components of laboratory equipment, so that the components can cooperate as a compact unified, fully automated system. In one embodiment, the invention provides a transporter for transporting specimen carriers to a processing station, comprising a first vertical input magazine of carriers, a second vertical output magazine of carriers, and a singulator located between the first and second magazines. The first and second magazines may be located one above the other. Preferably, the first and second magazines and the singulator are vertically aligned so that a carrier follows a straight vertical path. This provides a compact arrangement which takes up the minimum of floor space in the laboratory. The singulator receives carriers one-at-a-time from the input magazine and moves them vertically to a processing position. After processing, the singulator then delivers each carrier to an output location where it is transferred to the output magazine. The singulator may move a carrier upwards or downwards at the processing position independent of any movement of the carriers in the input and/or output magazines. Hence, the singulator can move a carrier to a predetermined distance from either the input magazine and/or the output magazine and can adjust this position in both directions. In particular, the singulator is adapted to move a carrier very accurately and finely about the processing position, e.g. to allow autofocusing. The sigulator may be described as micro-positioning, micro-stepping, micro-incrementing or micro-indexing. The drives for the input and output magazine are preferably simple gravity drives but the present invention is not limited thereto. The drive for the singulator is preferably a stepping motor or DC servo-motor. The processing position is at a location within the movement of the singulator.

The present invention may include in one embodiment a singulator comprising a release mechanism and a conveyor. The conveyor may comprise a pair of opposed belts, the belts having external protuberances or ledges. The belts are arranged vertically, so that the external ledges from the two belts define horizontally-oriented shelves which are capable of holding specimens such as carriers, slides, or other objects, in a generally horizontal orientation. The movement of the belts is preferably synchronized such that the specimens may be conveyed vertically up and down in their horizontal orientation while the belts move. The singulator is preferably adapted so that it not only transports the specimen carrier or slide to a viewing position but also

provides the necessary fine vertical movements for autofocusing at the viewing position. The singulator of the present invention may also be used to transport specimens to a processing position which does not involve optical scanning buut may

involve other processes.

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For this purpose, the transporter in accordance with the present invention may include a processing tool which may be moved horizontally to a location above and/or below the carrier at the processing position in the singulator. Further, it is preferred if the singulator spaces the selected carrier from the output and input magazines at the processing position to allow room for the processing tool to be introduced above and/or below the selected carrier. The processing tool is preferably an optical receiver for capturing an image, e.g. a video image of the contents of the carrier. The processing tool may be any other suitable tool, e.g. a microscope or a fluorescent microscope.

The present invention includes a method of transporting specimen carriers to a processing position, comprising the steps of: providing a vertical input magazine and a vertical output magazine of carriers; singling out a carrier from the input magazine and moving it vertically to the processing position which is located between the input and output magazines; and moving the carrier to an output location where it is transferred to the output magazine.

The step of moving the carrier to the processing position is preferably carried out in such a way that there is no simultaneous movement of the carriers in the input and/or output magazines. As only one carrier is moved to the processing position at a time, the load on a drive motor is low. Further, it is preferred if the singling out and moving steps space the selected carrier from the output and input magazines to allow room for a processing tool to be introduced above and/or below the selected carrier when the carrier is at the processing position. The present invention includes moving the processing tool horizontally to a position above and/or below the carrier at the processing position and withdrawing the processing tool after carrying out processing to allow the carriers to pass. The processing tool is preferably an optical receiver for capturing an image, e.g. a video image of the contents of the carrier. The processing tool may be any other suitable tool, e.g. a microscope or a fluorescent microscope.

The dependent claims each define a separate and individual embodiment of the present invention. The present invention will now be described with reference to the following drawings.

#### BRIEF DESCRIPTION OF THE DRAWINGS

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Fig. 1 is schematic side-view representation of the transport apparatus according to an embodiment of the present invention.

Fig. 2 is a schematic top view of the apparatus of Fig. 1.

Figure 3 is a detail of the cam mechanism for allowing a single specimen to enter the singulator.

Fig. 4 is a side view of one type of a specimen carrier which may be used with the present invention.

Fig. 5 is a schematic cross-section of an optical receiver in accordance with the present invention.

#### DETAILED DESCRIPTION OF THE ILLUSTRATIVE EMBODIMENTS

The present invention will be described with respect to certain embodiments and to certain schematic drawings but the present invention is not limited thereto but only by the claims.

Figs. 1 and 2 show schematically a side-view (with front plate 30 removed) and a top-view (with the input magazine 1 removed) of the transporter 100 in accordance with an embodiment of the present invention. Generally, the transporter 100 includes a vertical input magazine 1 of specimen carriers 8, a singulator 3 and an output magazine 6 of carriers 8. Each carrier 8 may contain one or more wells 9 in which liquid samples, in particular, cell cultures and other biological specimens may be placed. The word carrier is meant to be interpreted in a aide sense and may include petri dishes, trays, multiwell trays, microscope slides or holders for microscope slides or any other similar carrier, tray or specimen holder which casn be stacked in an input stack and moved to a processing position. The singulator 3 receives a single carrier 8' from the input magazine 1 via cam-driven release means 2. The singulator 3 then conveys the substantially horizontal carrier 8' vertically to a processing position 40. Singulator 3 can move the carrier 8' both upwards and downwards. Due to the fact that the singulator 3 only has to move a single carrier 8, the power required for the drive for the singulator 3 is low. Also, due to the low load on this drive, high positional accuracy and fine control of movement can be obtained, e.g. by using a stepper motor or DC servo-motor drive.

As shown schematically in Figs. 1 and 2, a singulator 3 according to an

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embodiment of the present invention includes a release mechanism 2 and a conveyor comprising four belts 16, 17 arranged in a generally vertical orientation. Each belt 16 runs on a pair of pulleys 25, 28. Each belt 17 runs on four pulleys 24, 26, 27, 29. Pulleys 24, 29 are mechanically biased, e.g. sprung loaded, and movable to stretch belt 17 in such a way that the carrier 8 is pressed against a guide plate 14 in the processing position 40. Belts 16, 17 have a plurality of protuberances or ledges 15, located on the outer surface of each belt and projecting substantially perpendicularly to the longitudinal direction of each belt. The belts 16, 17 are preferably made of rubber or other elastomer, or a rubber-like or elastomeric material, in particular fibre reinforced rubber or elastomer or rubber impregnated or elastomer impregnated textile or of some other flexible but preferably not stretchable material. The present invention is not limited to rubber or rubber substitutes. The portions of belts 16, 17 facing each other are spaced apart and move parallel to each other. The inner distance between the belts 16, 17 is greater than the outer dimension of the carriers 8, so as to allow the carriers 8 to fit between the belts 16, 17. The distance between the protuberances 15 on the two belts 16, 17 is less than the outer dimension of the carriers 8, so that each carrier 8 can be carried by four protuberances 8, one from each of the four belts 16, 17.

The pulleys 25, 28 of belt 16 rotate in a direction opposite to that of the pulleys 24, 26, 27, 29 of belt 17. The protuberances 15 on two belts 16, 17 which face each other move in the same vertical direction. The rotation of the pair of pulleys 25, 28 is preferably synchronized with the rotation of the four pulleys 25, 26, 27, 29 so that a carrier 8 supported by the promberances 15 remains in horizontal position during the movement. The carriers 8 may be moved either up or down by singulator 3, depending on the direction of rotation of the pulleys 24-29. Upper pulleys 27, 28 are each fixed to an axle 37, 38 respectively. Axles 37, 38 are journalled in two fixed, parallely spaced support plates 30. Axles 37, 38 are fixed at one end to pulleys 35, 36. Pulleys 35, 36 are driven by a further belt 11 which is driven by a motor 33 having a pulley 10 fixed to its output shaft. The belt 11 may be tensioned by a tensioning pulley 12 in a conventional way. Motor 33 is preferably a motor which can be driven accurately and in small increments, e.g. a stepping motor or a DC-servo-motor. Driving pulleys 35, 36 causes axles 37, 38 to rotate, hence driving pulleys 27, 28 and belts 16, 17. The backlash in the drive for belts 16, 17 should preferably be kept to a minimum by maintaining a constant positive tension in belts 16, 17 and/or to use a non-slip drive between the belts 16, 17

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and the pulleys 24, 26, 27, 29, e.g. driving belts and passsing profiled pulleys. In accordance with the present invention the backlash is preferably kept below 1 mm, more preferably 500 micron or less, and most preferably 300 micron or less. The backlash is preferably compensated for in the controller of the motor 33, e.g. on changes of direction the moteor is moved a set distance, ore than specified in order to take up the backlash. The repeatability of a vertical position, i.e. the accuracy of the system is preferably better than 100 micron, more preferably 50 micron or less and most preferably 30 micron or less. 10 micron is typical. An aspect of the present invention is the provision of what may be described as a micro-incrementing, micropositioning or micro-stepping singulator. This means that the singulator can be incremented in small steps of the order of 10 microns or less in one direction. One use of the singulator in accordance with the present invention is to bring a specimen carrier to a processing position and then to use the singulator to microindex the carrier up and down, e.g. to find a focussing position of a microscope. Thus, in accordance with the present invention, the singulator is not only a transporter for transporting a carrier from an input magazine to an output magazine, but also a singulator which can be halted at an intermediate position, moved accurately, if necessary, into a pre-determined vertical position and then can be incremented up and down about this position. All these movements can be carried out using the same motor and belt drive.

An operation of the transporter will now be described. A plurality of carriers 8 is first loaded into an input magazine 1. A weight 7 may be applied to the top of the magazine to provide a gravity drive therefor. Carriers 8 are prevented from entering the singulator 3 by a cam-driven release mechanism 2. Once allowed onto the conveyor 16,17, each carrier 8 is carried and conveyed by belts 16, 17. A carrier 8 is supported between protuberances 15 and is lowered to the processing position 40. By further movement of belts 16, 17 it is deposited into an output magazine 6.

To synchronize belts 16, 17 it is preferred if all of the pulleys 25, 26, 27, 28 have the same diameter, and are made to turn at the same rate. The pulleys 25-28 can be adjusted at the start so that the corresponding protuberances 15 on each belt combination 16, 17 are aligned horizontally. Sensor devices may be provided (not shown) to check that opposite protuberances 15 carry carriers 8 in a horizontal position. If a carrier 8 is not sufficiently horizontal, an error may cause an alarm.

The release mechanism 2 is driven in synchronism with the conveyor belts 16,

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17 so that only one carrier 8 is released from the input magazine 1 onto the conveyor belts 16, 17 at a time. Fig. 3 shows a detailed schematic representation of the cam 41 and the cam follower 45 of release mechanism 2. It is advantageous to drive the cam 41 with the same motor 33 as is used to drive axles 37, 38. By this means the required synchronism between the cam 41 and the conveyor belts 16, 17 can be maintained. As shown in Fig. 2, the cam 41 is fixed to axle 38 and is located between the pulleys 28. Two release mechanisms 2 are provided opposite each other for holding and releasing carriers 8. As shown the release mechanisms 2 are placed on either side of a carrier 8 close to a pulley 27, 28 respectively. The present invention is not limited thereto. For instance, the release mechanisms 2 may be placed across the dimension of the carrier 8 which is 90° to that shown in Fig. 2.

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A release mechanism 2 may include one or more (two are shown) support members 47 which are rotatably journalled on an axle 48. The release mechanism 2 also includes a cam follower 45 which bears on a cam 41 having one or more V-shaped notches 42. Each V-shaped notch 42 is bounded by sloping bearing surfaces 43, 44. Cam 41 may have a generally circular shape so that cam follower 45 remains at the same position as cam 41 rotates until a notch 42 is reached. As cam 41 rotates further cam follower 45 moves down the first sloping surface 43 and the support member 47 rotates about axle 48 making the gap between the ends of the support members 47 larger. This allows a carrier 8 to slide downwards past the ends of the support members 47. The position of the notch 42 is synchronized with the movement of belts 16, 17 so that a carrier 8' is lowered past the ends of support members 47 as protuberances 15 are at a suitable position to support this carrier 8'. As axle 38 continues to rotate the carrier 8' is lowered by movement of belts 16, 17. At the same time cam follower 45 rides up the surface 44 thus bring the end of support member 47 back to the sides of the next carrier 8 in the input magazine 1. Preferably, the relative position of the two sloping walls 43, 42 of a notch 42 may be varied so that the distance therebetween changes. For a given radius of cam 41, increasing the distance between the two sloping walls 43, 44 increases the distance that the sigulator 3 moves before the support member moves back up and supports the next carrier 8. Thus, by varying the distance between the walls 43, 44, the singulator 3 can be adapted to carriers 8 of different depths.

Preferably, a carrier 8 has the side-view form shown in Fig. 4 and in particular it is preferred if the carriers 8 are suitable for stacking, e.g. with a smaller upper part 51

than lower part 52. The next carrier 8 in the stack forms a lip 53. Preferably, carriers 8 are designed so that the top of one carrier seals off againts the bottom of the carrier above. This is advantageous when the carriers 8 contain liquids and evaportaion of these liquids should be prevented or reduced. As support member 47 returns to its initial position it engages with lip 53 of the next carrier 8 in the input magazine 1 and prevents this carrier from moving downwards any further at this time. The distance traveled by belts 16, 17 as cam 41 rotates from one notch 42 to the next is preferably equal to the travel of a carrier 8 from the top of the singulator to the bottom thereof, so that only one carrier 8 is present in the singulator 3 at any time. The present invention is not limited thereto. For instance two carriers 8 may be present in the singulator 3 at any one time provided these carriers 8 are adequately spaced apart.

Release of a carrier 8 from the singulator 3 to the output magazine 6 occurs automatically as belts 16, 17 travel around pulleys 25, 26. At this point the corresponding protuberances 15 move apart and allow each carrier 8 to descend into the output magazine 6. Output magazine 6 may include a plate 21 and flexible ropes 19 attached thereto. These ropes 19 run over the axles of the pulleys 25, 26 and are attached to flexible counter-weights 22 in housings 23. Movement of the ropes 19 may be enabled in one direction only by gravity and clamps 18 which press a rope 19 against the axle of the pulley 25 or 26 may prevent reverse motion. As the input magazine 6 moves down with increasing numbers of carriers 8, a longer length of counter-weight 22 is lifted up. By adjusting the weight of the flexible counter-weight 22, the additional weight of a carrier 8 in the output magazine 6 is compensated by the additional length of flexible counter-weight 22.

The transporter 100 described above is particularly useful for the delivery of carriers 8 to a optical viewing position 40 and for autofocus adjustments of the position of each carrier 8 at this position although the present invention is not limited only to optical viewing but includes additional or alternative operations such as laser cutting or trimming, addition of chemical compounds such as dies, solvents, reagents or similar, heating, cooling, drying, cutting etc. At the processing position 40 one or more tools 5 may be introduced horizontally from one or both sides of the carrier 8'. In addition tool or tools 5 may be introduced either above and/or below the carrier 8'. In particular, tool 5 may include an optical receiver 5A and a light source 5B arranged above and below the carrier 8' respectively. Optical receiver 5A and light source 5B may be attached to

an XY stage 4, which can move the tool 5 in a plane XY which is perpendicular to the movement direction of carrier 8' in the singulator 3 (the Z direction). Stage 4 may be driven in the orthogonal X and Y directions by motors 31, 32, e.g. stepping motors or DC-servo motors. In operation, the tool 5 may normally be kept retracted so that it does not interfere with the operation of the singulator 3 and carriers 8. Once a carrier 8 has reached the processing position 40, the tool 5A may be moved into the carrier region using the stage 4. The present invention is not limited to only one processing position 40 but may include several, whereby the contents of carrier 8 are processed sequentially at the processing positions.

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A purpose of the singulator 3 is to single out each carrier 8 to be measured and to provide sufficient distance between the singled out carrier 8' and the next carrier (stored at the bottom of the input magazine 1) so that the tool 5 may be moved into the space above and/or below the carrier 8' at the processing position 40. Preferably, sufficient space is also allowed at the processing position 40 between the singled out carrier 8' and the previous carrier 8 (stored at the top of the output magazine 6). Preferably, there is a free, unobstructed space above at least a central portion of the singled out carrier 8' which extends up to the bottom of the next carrier 8. The provision of this space is accomplished while keeping the input and output magazines 1, 6 very compact as each carrier 8 rests on the next one in these magazines. This free space is greater than the spacing of the carriers 8 in the input magazine 1 and may also be greater than the spacing between the protrusions or ledges 15 of the singulator 13. A further aspect of the singulator 3 is to provide vertical movements, both up and down, of the carrier 8' without any influence on the output magazine 6 or the input magazine 1. This allows Z direction movement of the carrier 8' to provide focusing of the contents of carrier 8' in the optical receiver 5A when the carrier is at the processing position 40.

An embodiment of an optical receiver 5A and light source 5B in accordance with the present invention is shown in schematic cross-section in Fig. 5. The optical receiver 5A may include a prism 5, at its outer end remote from a camera 34 to deflect light traveling vertically from the specimen carrier 8 along tube 52 in the horizontal direction towards a camera 34. Within the tube 52 of the optical receiver 5A one or more suitable filters, lenses 53 or diaphragms 54 may be placed to direct the light from the carrier 8 to the camera 34. Camera 34 may be a video camera attached to suitable

equipment for recording the video image of the specimen carrier 8, e.g. a personal computer (PC) with a video frame grabber and suitable digital storage means, e.g. a hard disk. Light for illuminating a carrier 8 may be optionally directed onto the specimen carrier 8' from above using a light source 5B including a fiber bundle 39 attached to the optical receiver 5A. Optical receiver 5A is preferably made telescopic so that its length can be adjusted to the focal length of the objective lens 53 used.

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Light source 5B may also include a prism at its extreme end to direct light from along the fibre optic cable 39 to the specimen carrier 8. Further, one or more suitable diaphragms 56, condensers 57 or filters 58 may be placed between the prism 55 and the fibre optic cable 39 to direct light travelling along the fibre optic cable 39 to the prism and to control its projected spot size and colour.

Instead of being attached to an XY stage, the optical receiver 5A may be attached to an XYZ stage (not shown) so that focusing of the image received by the camera 34 may be accomplished by Z direction movements of the stage. However, it is preferred in accordance with the present invention if the Z direction movements are carried out by the singulator 3. The reason for this is that the optical receiver 5A and camera 34 are heavy and movements in particular acceleration of these parts can cause vibrations and shocks to the complete system which can disturb the focusing operations and slow these down as it is necessary to wait between each movement until vibrations have ceased in order to capture an image accurately. The carrier 8 and the conveyor belts 16, 17 are light and cause little mechanical disturbance thus allowing much more rapid focusing onto the contents of carrier 8. This accelerates automatic screening.

To improve focusing still further, a device 20 may be provided for clamping the carrier 8' at the focused position 40. Clamping device 20 (best shown in Fig. 1) may include two pulleys 24, 29 for engagement with the outer portion and inner portion of belt 17 respectively. Between the two pulleys 24, 29 is placed a small actuator which drives pulley 29 against the side of carrier 8' when suitably stimulated. The actuator may be a spring. By this means the carrier 8' is clamped between the pulley 29 and a fixed plate 14 in a steady position even when optical receiver 5A is moved sideways by stage 4, e.g. when moving from one field to another or from one well to another of the specimen carrier 8'. A position sensor 13 such as a microswitch may be used to indicate when the carrier 8 is at the processing position 40. The output of the sensor 13 may be used for activating the movement of tool 5 into the processing area.

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The control of the various components of transporter 100 including the movements of stage 4, belts 16 and 17 and clamp 20 may be carried out using suitable 3-axis control devices, e.g. MultiControl 2000 supplied by Märzhauzer and a PC computer running suitable programmable software.

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While the invention has been shown and described with reference to preferred embodiments, it will be understood by those skilled in the art that various changes or modifications in form and detail may be made without departing from the scope and spirit of this invention as defined in the attached claims. For example, the input magazine 1 has been described above and shown in the attached drawings as being located above the output magazine but the present invention is not limited thereto. The input magazine may be placed below the output magazine with the singulator between the two. In this case the carriers in the input magazine are driven upwards and the singulator receives a carrier and conveys it upwards towards the processing position and the output magazine. This embodiment of the present invention requires motor drives for the input and output magazines which is less preferred. Alternatively, in accordance with an embodiment of the present invention devices may be included for transporting the carrier 8 when it has arrived at the processing poistion 40 to a different processing area, e.g. by moving the carrier 8 horizontally out of the singulator 3 and optionally returing it to the singulator 3 after this processing.

The present invention has been described with reference to an input magazine and an output magazine of carriers but the present invention is not limited thereto. The carrier may be moved to the processing position and removed therefrom to some other place so that only the input magazine is generally required for the present invention. Alternatively, the carriers may no longer be required after processing at the processing position and the carrier may simply be discharged from the bottom of the singulator. For instance, the carrier may simply fall from the bottom of the singulator into a bin.

#### CLAIMS

1. A transporter for transporting carriers to a processing station, comprising:
a first vertical input magazine of carriers, a second output magazine of carriers, the first
and second magazines being located one above the other in either order; and
a singulator located between the output of the first magazine and the input of the second
magazine, the singulator being adapted to receive one carrier at a time from the first
magazine and to move the carrier from the first magazine vertically to a position
intermediate the first and the second magazines and to halt the carrier at this position.

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2. A transporter for transporting specimen carriers, comprising:
a first vertical input magazine of carriers, the carriers in the first magazine being spaced at
a carrier spacing distance; a singulator adapted to receive one carrier at a time from the
first magazine and to move a received carrier vertically from a first position at the output
of the first magazine to a second position at which a carrier may be discharged from the
singulator; wherein the singulator is adapted to move a received carrier and to halt it at an
intermediate position located between the first and second positions to space the received
carrier from the next carrier in the first input magazine by a distance greater than the
carrier spacing distance.

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3. The transporter according to claim 1 or 2, wherein the singulator is adapted to move a received carrier incrementally upwards or downwards at the intermediate position independent of any movement of the carriers in the first and/or second magazines.

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4. The transporter according to any previous claim wherein the singulator comprises a release mechanism and a conveyor, the release mechanism releasing carriers from the first magazine one-at-a-time to the conveyor.

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5. The transporter according to any of the previous claims wherein the conveyor of the singulator comprises a pair of opposed belts, the belts having external protuberances or ledges and the belts being arranged vertically, so that the external ledges from the two belts define horizontally-oriented shelves which are capable of holding a received carrier in a substantially horizontal orientation.

6. The transporter according to any previous claim, further comprising a processing tool movable horizontally to a location above and/or below the carrier when the carrier is at the intermediate position in the singulator.

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- 7. The transporter according to claim 6, wherein the processing tool is an optical receiver for capturing an image, in particular a video image of the contents of the carrier.
- 8. The transporter according to claim 7, wherein the optical receiver comprises a microscope.
  - 9. The transporter according to any previous claim, wherein the carriers are Petri dishes, multi-well plates or any other device containing biological specimens.
- 15 10. A method of transporting specimen carriers to a processing position, comprising the steps of:

providing a vertical input magazine and an output magazine of carriers, one above the other;

singling out a carrier from the input magazine and moving it vertically to an intermediate position which is located between the input and output magazines and halting it there; and moving the carrier to an output location where it is transferred to the output magazine.

11. The method according to claim 10, further comprising moving the carrier incrementally up or down at the intermediate position using the singulator.

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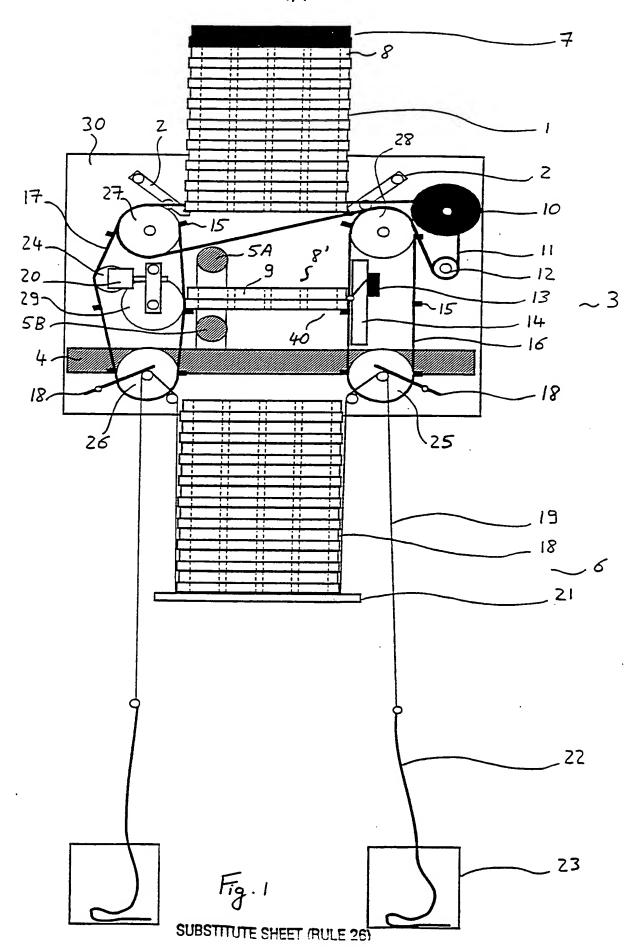
12. The method according to claim 10 or 11, further comprising the steps of: moving a processing tool horizontally to a position above and/or below the carrier at the intermediate position; and withdrawing the processing tool after carrying out processing.

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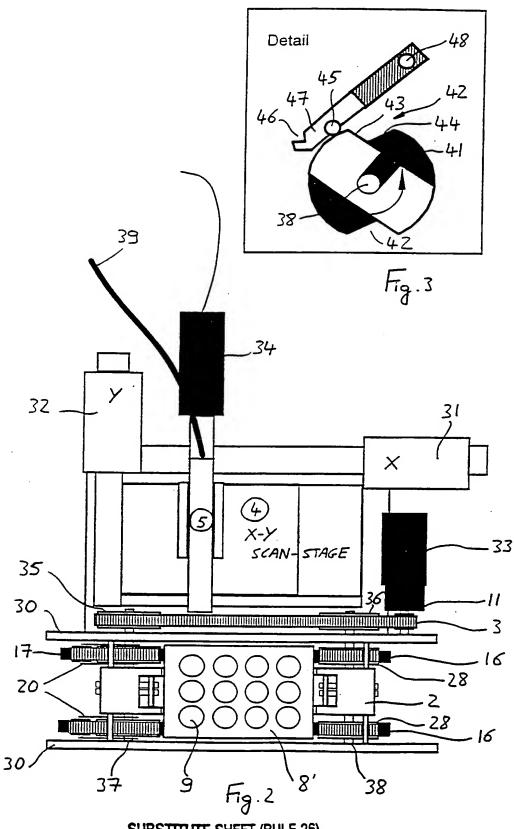
13. The method according to any of claims 10 to 12, wherein the movement of the carrier to the intermediate position is independent of movements of the carriers in the input and output magazines

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14. The method according to claim 12 or 13, wherein the processing tool is an optical receiver.

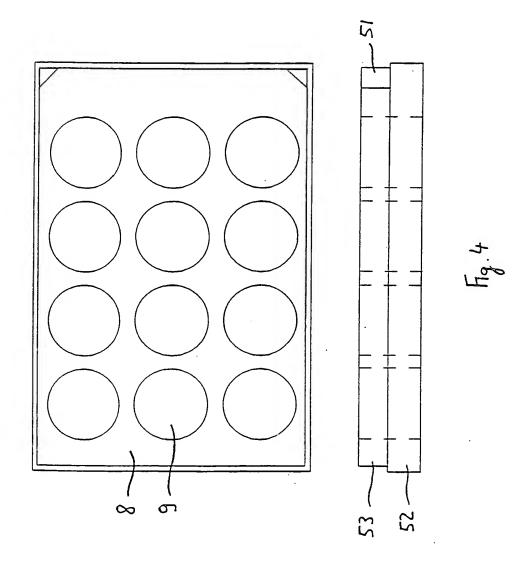


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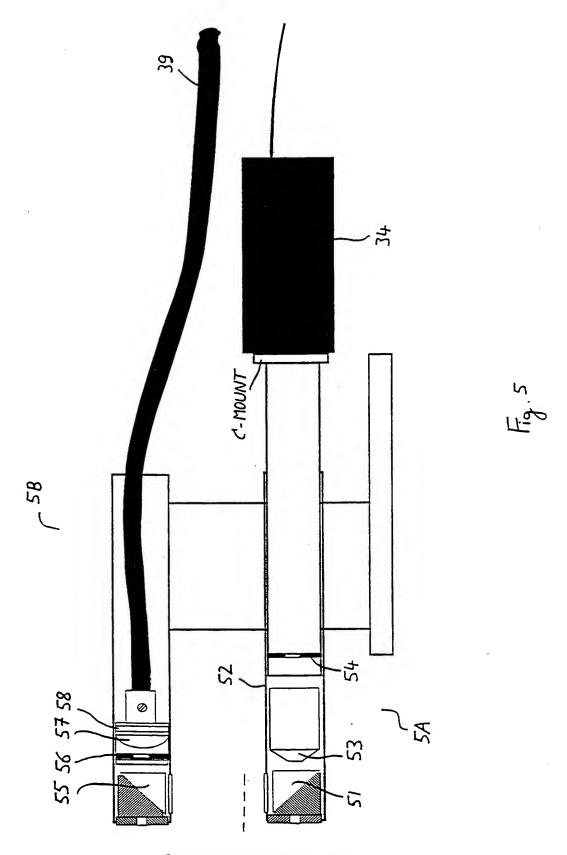


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# APPARATUS FOR TRANSPORTING CARRIERS TO A PROCESSING STATION AND METHOD OF OPERATING THE SAME

The present invention relates to handling of objects, particularly objects to be scanned optically or viewed as for instance microscope slides, specimen carriers, trays or Petri dishes. In particular, the present invention relates to an apparatus and method for transporting carriers, such as microscope slides or trays to a processing position.

#### TECHNICAL BACKGROUND OF THE INVENTION

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Devices for conveying specimens particularly for clinical laboratories are known from DE-U-297 12 535, WO 97/39348, WO 93/06516. However, most of these devices need to grasp the specimen, or push the specimen sideways out of a vertical magazine to a viewing position or need to move them in some other way to a viewing or processing position. It is often inconvenient grasp or push a delicate specimen holder such as a microscope slide, either because the slide is fragile, or because the specimen, or the indicia imprinted on the slide, can easily be abraded and thus obliterated. In addition, the known devices require a separate mechanism for autofocusing onto the specimen to be viewed. For example, autofocusing mechanisms are known from WO 96/01438, US 3,721,759, WO 96/10196, US 5,790,692, US 5,790,710. These require a stage for moving the specimen in at least two dimensions and often in three. This stage is in addition to the vertical delivery systems described above. Each specimen must be conveyed to a first position with one mechanism, transferred to a viewing position with another mechanism and then autofocused using yet another mechanism. This makes the combination expensive, bulky and complicated.

It is an object of the present invention to provide a compact system and method for delivery of specimens, particularly those to be viewed optically, to a processing or viewing position.

It is a further object of the present invention to provide an apparatus and method for obtaining an autofocus position for an optical instrument in a short time.

It is a further object of the present invention to provide an apparatus and method for obtaining an autofocus position for an optical instrument which is simple, compact and less expensive than known apparatus and methods.

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## SUMMARY OF THE INVENTION

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One aspect of the present invention is to integrate components of laboratory equipment, so that the components can cooperate as a compact unified, fully automated system. In one embodiment, the invention provides a transporter for transporting specimen carriers to a processing station, comprising a first vertical input magazine of carriers, a second vertical output magazine of carriers, and a singulator located between the first and second magazines. The first and second magazines may be located one above the other. Preferably, the first and second magazines and the singulator are vertically aligned so that a carrier follows a straight vertical path. This provides a compact arrangement which takes up the minimum of floor space in the laboratory. The singulator receives carriers one-at-a-time from the input magazine and moves them vertically to a processing position. After processing, the singulator then delivers each carrier to an output location where it is transferred to the output magazine. The singulator may move a carrier upwards or downwards at the processing position independent of any movement of the carriers in the input and/or output magazines. Hence, the singulator can move a carrier to a predetermined distance from either the input magazine and/or the output magazine and can adjust this position in both directions. In particular, the singulator is adapted to move a carrier very accurately and finely about the processing position, e.g. to allow autofocusing. The sigulator may be described as micro-positioning, micro-stepping, micro-incrementing or micro-indexing. The drives for the input and output magazine are preferably simple gravity drives but the present invention is not limited thereto. The drive for the singulator is preferably a stepping motor or DC servo-motor. The processing position is at a location within the movement of the singulator.

The present invention may include in one embodiment a singulator comprising a release mechanism and a conveyor. The conveyor may comprise a pair of opposed belts, the belts having external protuberances or ledges. The belts are arranged vertically, so that the external ledges from the two belts define horizontally-oriented shelves which are capable of holding specimens such as carriers, slides, or other objects, in a generally horizontal orientation. The movement of the belts is preferably synchronized such that the specimens may be conveyed vertically up and down in their horizontal orientation while the belts move. The singulator is preferably adapted so that it not only transports the specimen carrier or slide to a viewing position but also

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provides the necessary fine vertical movements for autofocusing at the viewing position. The singulator of the present invention may also be used to transport specimens to a processing position which does not involve optical scanning buut may involve other processes.

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For this purpose, the transporter in accordance with the present invention may include a processing tool which may be moved horizontally to a location above and/or below the carrier at the processing position in the singulator. Further, it is preferred if the singulator spaces the selected carrier from the output and input magazines at the processing position to allow room for the processing tool to be introduced above and/or below the selected carrier. The processing tool is preferably an optical receiver for capturing an image, e.g. a video image of the contents of the carrier. The processing tool may be any other suitable tool, e.g. a microscope or a fluorescent microscope.

The present invention includes a method of transporting specimen carriers to a processing position, comprising the steps of: providing a vertical input magazine and a vertical output magazine of carriers; singling out a carrier from the input magazine and moving it vertically to the processing position which is located between the input and output magazines; and moving the carrier to an output location where it is transferred to the output magazine.

The step of moving the carrier to the processing position is preferably carried out in such a way that there is no simultaneous movement of the carriers in the input and/or output magazines. As only one carrier is moved to the processing position at a time, the load on a drive motor is low. Further, it is preferred if the singling out and moving steps space the selected carrier from the output and input magazines to allow room for a processing tool to be introduced above and/or below the selected carrier when the carrier is at the processing position. The present invention includes moving the processing tool horizontally to a position above and/or below the carrier at the processing position and withdrawing the processing tool after carrying out processing to allow the carriers to pass. The processing tool is preferably an optical receiver for capturing an image, e.g. a video image of the contents of the carrier. The processing tool may be any other suitable tool, e.g. a microscope or a fluorescent microscope.

The dependent claims each define a separate and individual embodiment of the present invention. The present invention will now be described with reference to the following drawings.

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#### BRIEF DESCRIPTION OF THE DRAWINGS

Fig. 1 is schematic side-view representation of the transport apparatus according to an embodiment of the present invention.

Fig. 2 is a schematic top view of the apparatus of Fig. 1.

Figure 3 is a detail of the cam mechanism for allowing a single specimen to enter the singulator.

Fig. 4 is a side view of one type of a specimen carrier which may be used with the present invention.

Fig. 5 is a schematic cross-section of an optical receiver in accordance with the present invention.

#### DETAILED DESCRIPTION OF THE ILLUSTRATIVE EMBODIMENTS

The present invention will be described with respect to certain embodiments and to certain schematic drawings but the present invention is not limited thereto but only by the claims.

Figs. 1 and 2 show schematically a side-view (with front plate 30 removed) and a top-view (with the input magazine 1 removed) of the transporter 100 in accordance with an embodiment of the present invention. Generally, the transporter 100 includes a vertical input magazine 1 of specimen carriers 8, a singulator 3 and an output magazine 6 of carriers 8. Each carrier 8 may contain one or more wells 9 in which liquid samples, in particular, cell cultures and other biological specimens may be placed. The word carrier is meant to be interpreted in a aide sense and may include petri dishes, trays, multiwell trays, microscope slides or holders for microscope slides or any other similar carrier, tray or specimen holder which casn be stacked in an input stack and moved to a processing position. The singulator 3 receives a single carrier 8' from the input magazine 1 via cam-driven release means 2. The singulator 3 then conveys the substantially horizontal carrier 8' vertically to a processing position 40. Singulator 3 can move the carrier 8' both upwards and downwards. Due to the fact that the singulator 3 only has to move a single carrier 8, the power required for the drive for the singulator 3 is low. Also, due to the low load on this drive, high positional accuracy and fine control of movement can be obtained, e.g. by using a stepper motor or DC servo-motor drive.

As shown schematically in Figs. 1 and 2, a singulator 3 according to an

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embodiment of the present invention includes a release mechanism 2 and a conveyor comprising four belts 16, 17 arranged in a generally vertical orientation. Each belt 16 runs on a pair of pulleys 25, 28. Each belt 17 runs on four pulleys 24, 26, 27, 29. Pulleys 24, 29 are mechanically biased, e.g. sprung loaded, and movable to stretch belt 17 in such a way that the carrier 8 is pressed against a guide plate 14 in the processing position 40. Belts 16, 17 have a plurality of protuberances or ledges 15, located on the outer surface of each belt and projecting substantially perpendicularly to the longitudinal direction of each belt. The belts 16, 17 are preferably made of rubber or other elastomer, or a rubber-like or elastomeric material, in particular fibre reinforced rubber or elastomer or rubber impregnated or elastomer impregnated textile or of some other flexible but preferably not stretchable material. The present invention is not limited to rubber or rubber substitutes. The portions of belts 16, 17 facing each other are spaced apart and move parallel to each other. The inner distance between the belts 16, 17 is greater than the outer dimension of the carriers 8, so as to allow the carriers 8 to fit between the belts 16, 17. The distance between the protuberances 15 on the two belts 16, 17 is less than the outer dimension of the carriers 8, so that each carrier 8 can be carried by four protuberances 8, one from each of the four belts 16, 17.

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The pulleys 25, 28 of belt 16 rotate in a direction opposite to that of the pulleys 24, 26, 27, 29 of belt 17. The protuberances 15 on two belts 16, 17 which face each other move in the same vertical direction. The rotation of the pair of pulleys 25, 28 is preferably synchronized with the rotation of the four pulleys 25, 26, 27, 29 so that a carrier 8 supported by the protuberances 15 remains in horizontal position during the movement. The carriers 8 may be moved either up or down by singulator 3, depending on the direction of rotation of the pulleys 24-29. Upper pulleys 27, 28 are each fixed to an axle 37, 38 respectively. Axles 37, 38 are journalled in two fixed, parallely spaced support plates 30. Axles 37, 38 are fixed at one end to pulleys 35, 36. Pulleys 35, 36 are driven by a further belt 11 which is driven by a motor 33 having a pulley 10 fixed to its output shaft. The belt 11 may be tensioned by a tensioning pulley 12 in a conventional way. Motor 33 is preferably a motor which can be driven accurately and in small increments, e.g. a stepping motor or a DC-servo-motor. Driving pulleys 35, 36 causes axles 37, 38 to rotate, hence driving pulleys 27, 28 and belts 16, 17. The backlash in the drive for belts 16, 17 should preferably be kept to a minimum by maintaining a constant positive tension in belts 16, 17 and/or to use a non-slip drive between the belts 16, 17

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and the pulleys 24, 26, 27, 29, e.g. driving belts and passsing profiled pulleys. In accordance with the present invention the backlash is preferably kept below 1 mm, more preferably 500 micron or less, and most preferably 300 micron or less. The backlash is preferably compensated for in the controller of the motor 33, e.g. on changes of direction the moteor is moved a set distance, ore than specified in order to take up the backlash. The repeatability of a vertical position, i.e. the accuracy of the system is preferably better than 100 micron, more preferably 50 micron or less and most preferably 30 micron or less. 10 micron is typical. An aspect of the present invention is the provision of what may be described as a micro-incrementing, micropositioning or micro-stepping singulator. This means that the singulator can be incremented in small steps of the order of 10 microns or less in one direction. One use of the singulator in accordance with the present invention is to bring a specimen carrier to a processing position and then to use the singulator to microindex the carrier up and down, e.g. to find a focussing position of a microscope. Thus, in accordance with the present invention, the singulator is not only a transporter for transporting a carrier from an input magazine to an output magazine, but also a singulator which can be halted at an intermediate position, moved accurately, if necessary, into a pre-determined vertical position and then can be incremented up and down about this position. All these movements can be carried out using the same motor and belt drive.

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An operation of the transporter will now be described. A plurality of carriers 8 is first loaded into an input magazine 1. A weight 7 may be applied to the top of the magazine to provide a gravity drive therefor. Carriers 8 are prevented from entering the singulator 3 by a cam-driven release mechanism 2. Once allowed onto the conveyor 16,17, each carrier 8 is carried and conveyed by belts 16, 17. A carrier 8 is supported between protuberances 15 and is lowered to the processing position 40. By further movement of belts 16, 17 it is deposited into an output magazine 6.

To synchronize belts 16, 17 it is preferred if all of the pulleys 25, 26, 27, 28 have the same diameter, and are made to turn at the same rate. The pulleys 25-28 can be adjusted at the start so that the corresponding protuberances 15 on each belt combination 16, 17 are aligned horizontally. Sensor devices may be provided (not shown) to check that opposite protuberances 15 carry carriers 8 in a horizontal position. If a carrier 8 is not sufficiently horizontal, an error may cause an alarm.

The release mechanism 2 is driven in synchronism with the conveyor belts 16,

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17 so that only one carrier 8 is released from the input magazine 1 onto the conveyor belts 16, 17 at a time. Fig. 3 shows a detailed schematic representation of the cam 41 and the cam follower 45 of release mechanism 2. It is advantageous to drive the cam 41 with the same motor 33 as is used to drive axles 37, 38. By this means the required synchronism between the cam 41 and the conveyor belts 16, 17 can be maintained. As shown in Fig. 2, the cam 41 is fixed to axle 38 and is located between the pulleys 28. Two release mechanisms 2 are provided opposite each other for holding and releasing carriers 8. As shown the release mechanisms 2 are placed on either side of a carrier 8 close to a pulley 27, 28 respectively. The present invention is not limited thereto. For instance, the release mechanisms 2 may be placed across the dimension of the carrier 8 which is 90° to that shown in Fig. 2.

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A release mechanism 2 may include one or more (two are shown) support members 47 which are rotatably journalled on an axle 48. The release mechanism 2 also includes a cam follower 45 which bears on a cam 41 having one or more V-shaped notches 42. Each V-shaped notch 42 is bounded by sloping bearing surfaces 43, 44. Cam 41 may have a generally circular shape so that cam follower 45 remains at the same position as cam 41 rotates until a notch 42 is reached. As cam 41 rotates further cam follower 45 moves down the first sloping surface 43 and the support member 47 rotates about axle 48 making the gap between the ends of the support members 47 larger. This allows a carrier 8 to slide downwards past the ends of the support members 47. The position of the notch 42 is synchronized with the movement of belts 16, 17 so that a carrier 8' is lowered past the ends of support members 47 as protuberances 15 are at a suitable position to support this carrier 8'. As axle 38 continues to rotate the carrier 8' is lowered by movement of belts 16, 17. At the same time cam follower 45 rides up the surface 44 thus bring the end of support member 47 back to the sides of the next carrier 8 in the input magazine 1. Preferably, the relative position of the two sloping walls 43, 42 of a notch 42 may be varied so that the distance therebetween changes. For a given radius of cam 41, increasing the distance between the two sloping walls 43, 44 increases the distance that the sigulator 3 moves before the support member moves back up and supports the next carrier 8. Thus, by varying the distance between the walls 43, 44, the singulator 3 can be adapted to carriers 8 of different depths.

Preferably, a carrier 8 has the side-view form shown in Fig. 4 and in particular it is preferred if the carriers 8 are suitable for stacking, e.g. with a smaller upper part 51

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than lower part 52. The next carrier 8 in the stack forms a lip 53. Preferably, carriers 8 are designed so that the top of one carrier seals off againts the bottom of the carrier above. This is advantageous when the carriers 8 contain liquids and evaportaion of these liquids should be prevented or reduced. As support member 47 returns to its initial position it engages with lip 53 of the next carrier 8 in the input magazine 1 and prevents this carrier from moving downwards any further at this time. The distance traveled by belts 16, 17 as cam 41 rotates from one notch 42 to the next is preferably equal to the travel of a carrier 8 from the top of the singulator to the bottom thereof, so that only one carrier 8 is present in the singulator 3 at any time. The present invention is not limited thereto. For instance two carriers 8 may be present in the singulator 3 at any one time provided these carriers 8 are adequately spaced apart.

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Release of a carrier 8 from the singulator 3 to the output magazine 6 occurs automatically as belts 16, 17 travel around pulleys 25, 26. At this point the corresponding protuberances 15 move apart and allow each carrier 8 to descend into the output magazine 6. Output magazine 6 may include a plate 21 and flexible ropes 19 attached thereto. These ropes 19 run over the axles of the pulleys 25, 26 and are attached to flexible counter-weights 22 in housings 23. Movement of the ropes 19 may be enabled in one direction only by gravity and clamps 18 which press a rope 19 against the axle of the pulley 25 or 26 may prevent reverse motion. As the input magazine 6 moves down with increasing numbers of carriers 8, a longer length of counter-weight 22 is lifted up. By adjusting the weight of the flexible counter-weight 22, the additional weight of a carrier 8 in the output magazine 6 is compensated by the additional length of flexible counter-weight 22.

The transporter 100 described above is particularly useful for the delivery of carriers 8 to a optical viewing position 40 and for autofocus adjustments of the position of each carrier 8 at this position although the present invention is not limited only to optical viewing but includes additional or alternative operations such as laser cutting or trimming, addition of chemical compounds such as dies, solvents, reagents or similar, heating, cooling, drying, cutting etc. At the processing position 40 one or more tools 5 may be introduced horizontally from one or both sides of the carrier 8'. In addition tool or tools 5 may be introduced either above and/or below the carrier 8'. In particular, tool 5 may include an optical receiver 5A and a light source 5B arranged above and below the carrier 8' respectively. Optical receiver 5A and light source 5B may be attached to

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an XY stage 4, which can move the tool 5 in a plane XY which is perpendicular to the movement direction of carrier 8' in the singulator 3 (the Z direction). Stage 4 may be driven in the orthogonal X and Y directions by motors 31, 32, e.g. stepping motors or DC-servo motors. In operation, the tool 5 may normally be kept retracted so that it does not interfere with the operation of the singulator 3 and carriers 8. Once a carrier 8 has reached the processing position 40, the tool 5A may be moved into the carrier region using the stage 4. The present invention is not limited to only one processing position 40 but may include several, whereby the contents of carrier 8 are processed sequentially at the processing positions.

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A purpose of the singulator 3 is to single out each carrier 8 to be measured and to provide sufficient distance between the singled out carrier 8' and the next carrier (stored at the bottom of the input magazine 1) so that the tool 5 may be moved into the space above and/or below the carrier 8' at the processing position 40. Preferably, sufficient space is also allowed at the processing position 40 between the singled out carrier 8' and the previous carrier 8 (stored at the top of the output magazine 6). Preferably, there is a free, unobstructed space above at least a central portion of the singled out carrier 8' which extends up to the bottom of the next carrier 8. The provision of this space is accomplished while keeping the input and output magazines 1, 6 very compact as each carrier 8 rests on the next one in these magazines. This free space is greater than the spacing of the carriers 8 in the input magazine 1 and may also be greater than the spacing between the protrusions or ledges 15 of the singulator 13. A further aspect of the singulator 3 is to provide vertical movements, both up and down, of the carrier 8' without any influence on the output magazine 6 or the input magazine 1. This allows Z direction movement of the carrier 8' to provide focusing of the contents of carrier 8' in the optical receiver 5A when the carrier is at the processing position 40.

An embodiment of an optical receiver 5A and light source 5B in accordance with the present invention is shown in schematic cross-section in Fig. 5. The optical receiver 5A may include a prism 5, at its outer end remote from a camera 34 to deflect light traveling vertically from the specimen carrier 8 along tube 52 in the horizontal direction towards a camera 34. Within the tube 52 of the optical receiver 5A one or more suitable filters, lenses 53 or diaphragms 54 may be placed to direct the light from the carrier 8 to the camera 34. Camera 34 may be a video camera attached to suitable

equipment for recording the video image of the specimen carrier 8, e.g. a personal computer (PC) with a video frame grabber and suitable digital storage means, e.g. a hard disk. Light for illuminating a carrier 8 may be optionally directed onto the specimen carrier 8' from above using a light source 5B including a fiber bundle 39 attached to the optical receiver 5A. Optical receiver 5A is preferably made telescopic so that its length can be adjusted to the focal length of the objective lens 53 used.

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Light source 5B may also include a prism at its extreme end to direct light from along the fibre optic cable 39 to the specimen carrier 8. Further, one or more suitable diaphragms 56, condensers 57 or filters 58 may be placed between the prism 55 and the fibre optic cable 39 to direct light travelling along the fibre optic cable 39 to the prism and to control its projected spot size and colour.

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Instead of being attached to an XY stage, the optical receiver 5A may be attached to an XYZ stage (not shown) so that focusing of the image received by the camera 34 may be accomplished by Z direction movements of the stage. However, it is preferred in accordance with the present invention if the Z direction movements are carried out by the singulator 3. The reason for this is that the optical receiver 5A and camera 34 are heavy and movements in particular acceleration of these parts can cause vibrations and shocks to the complete system which can disturb the focusing operations and slow these down as it is necessary to wait between each movement until vibrations have ceased in order to capture an image accurately. The carrier 8 and the conveyor belts 16, 17 are light and cause little mechanical disturbance thus allowing much more rapid focusing onto the contents of carrier 8. This accelerates automatic screening.

To improve focusing still further, a device 20 may be provided for clamping the carrier 8' at the focused position 40. Clamping device 20 (best shown in Fig. 1) may include two pulleys 24, 29 for engagement with the outer portion and inner portion of belt 17 respectively. Between the two pulleys 24, 29 is placed a small actuator which drives pulley 29 against the side of carrier 8' when suitably stimulated. The actuator may be a spring. By this means the carrier 8' is clamped between the pulley 29 and a fixed plate 14 in a steady position even when optical receiver 5A is moved sideways by stage 4, e.g. when moving from one field to another or from one well to another of the specimen carrier 8'. A position sensor 13 such as a microswitch may be used to indicate when the carrier 8 is at the processing position 40. The output of the sensor 13 may be used for activating the movement of tool 5 into the processing area.

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The control of the various components of transporter 100 including the movements of stage 4, belts 16 and 17 and clamp 20 may be carried out using suitable 3-axis control devices, e.g. MultiControl 2000 supplied by Märzhauzer and a PC computer running suitable programmable software.

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While the invention has been shown and described with reference to preferred embodiments, it will be understood by those skilled in the art that various changes or modifications in form and detail may be made without departing from the scope and spirit of this invention as defined in the attached claims. For example, the input magazine 1 has been described above and shown in the attached drawings as being located above the output magazine but the present invention is not limited thereto. The input magazine may be placed below the output magazine with the singulator between the two. In this case the carriers in the input magazine are driven upwards and the singulator receives a carrier and conveys it upwards towards the processing position and the output magazine. This embodiment of the present invention requires motor drives for the input and output magazines which is less preferred. Alternatively, in accordance with an embodiment of the present invention devices may be included for transporting the carrier 8 when it has arrived at the processing poistion 40 to a different processing area, e.g. by moving the carrier 8 horizontally out of the singulator 3 and optionally returing it to the singulator 3 after this processing.

The present invention has been described with reference to an input magazine and an output magazine of carriers but the present invention is not limited thereto. The carrier may be moved to the processing position and removed therefrom to some other place so that only the input magazine is generally required for the present invention. Alternatively, the carriers may no longer be required after processing at the processing position and the carrier may simply be discharged from the bottom of the singulator. For instance, the carrier may simply fall from the bottom of the singulator into a bin.

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## **CLAIMS**

1. A transporter for transporting carriers to a processing station, comprising:
a first vertical input magazine of carriers, a second output magazine of carriers, the first
and second magazines being located one above the other in either order; and
a singulator located between the output of the first magazine and the input of the second
magazine, the singulator being adapted to receive one carrier at a time from the first
magazine and to move the carrier from the first magazine vertically to a position
intermediate the first and the second magazines and to halt the carrier at this position.

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- 2. A transporter for transporting specimen carriers, comprising:
  a first vertical input magazine of carriers, the carriers in the first magazine being spaced at
  a carrier spacing distance; a singulator adapted to receive one carrier at a time from the
  first magazine and to move a received carrier vertically from a first position at the output
  of the first magazine to a second position at which a carrier may be discharged from the
  singulator; wherein the singulator is adapted to move a received carrier and to halt it at an
  intermediate position located between the first and second positions to space the received
  carrier from the next carrier in the first input magazine by a distance greater than the
  carrier spacing distance.

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- 3. The transporter according to claim 1 or 2, wherein the singulator is adapted to move a received carrier incrementally upwards or downwards at the intermediate position independent of any movement of the carriers in the first and/or second magazines.
- 4. The transporter according to any previous claim wherein the singulator comprises a release mechanism and a conveyor, the release mechanism releasing carriers from the first magazine one-at-a-time to the conveyor.
- 5. The transporter according to any of the previous claims wherein the conveyor of the singulator comprises a pair of opposed belts, the belts having external protuberances or ledges and the belts being arranged vertically, so that the external ledges from the two belts define horizontally-oriented shelves which are capable of holding a received carrier in a substantially horizontal orientation.

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6. The transporter according to any previous claim, further comprising a processing tool movable horizontally to a location above and/or below the carrier when the carrier is at the intermediate position in the singulator.

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- 7. The transporter according to claim 6, wherein the processing tool is an optical receiver for capturing an image, in particular a video image of the contents of the carrier.
- 8. The transporter according to claim 7, wherein the optical receiver comprises a microscope.
  - 9. The transporter according to any previous claim, wherein the carriers are Petri dishes, multi-well plates or any other device containing biological specimens.
- 10. A method of transporting specimen carriers to a processing position, comprising the steps of:
  - providing a vertical input magazine and an output magazine of carriers, one above the other;
  - singling out a carrier from the input magazine and moving it vertically to an intermediate position which is located between the input and output magazines and halting it there; and moving the carrier to an output location where it is transferred to the output magazine.
    - 11. The method according to claim 10, further comprising moving the carrier incrementally up or down at the intermediate position using the singulator.

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12. The method according to claim 10 or 11, further comprising the steps of: moving a processing tool horizontally to a position above and/or below the carrier at the intermediate position; and withdrawing the processing tool after carrying out processing.

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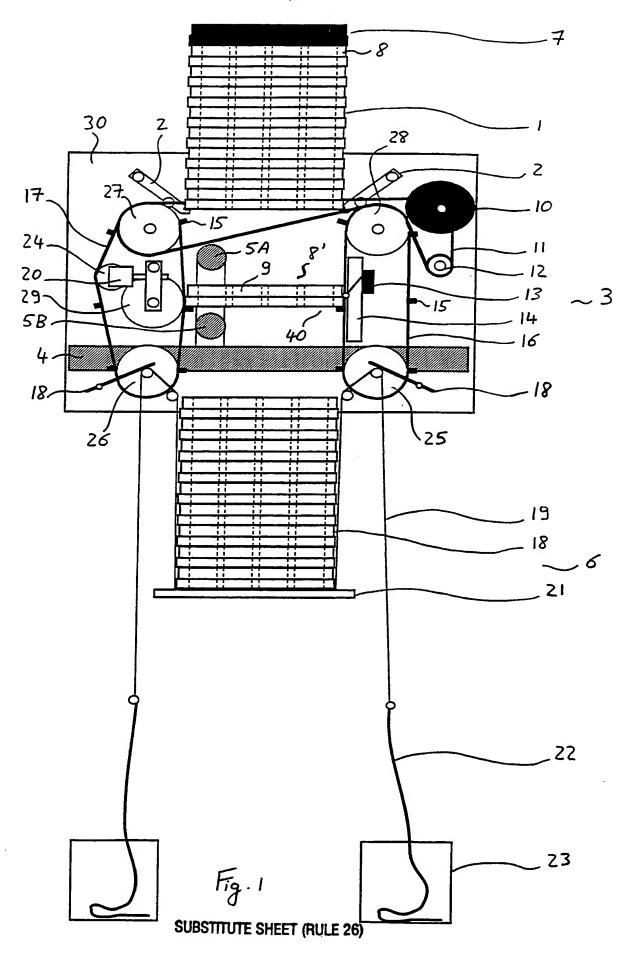
13. The method according to any of claims 10 to 12, wherein the movement of the carrier to the intermediate position is independent of movements of the carriers in the input and output magazines

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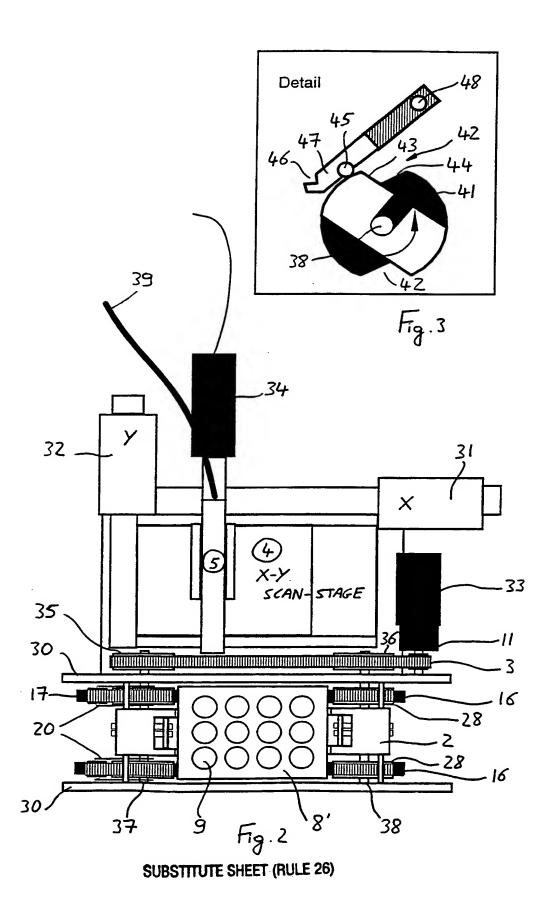
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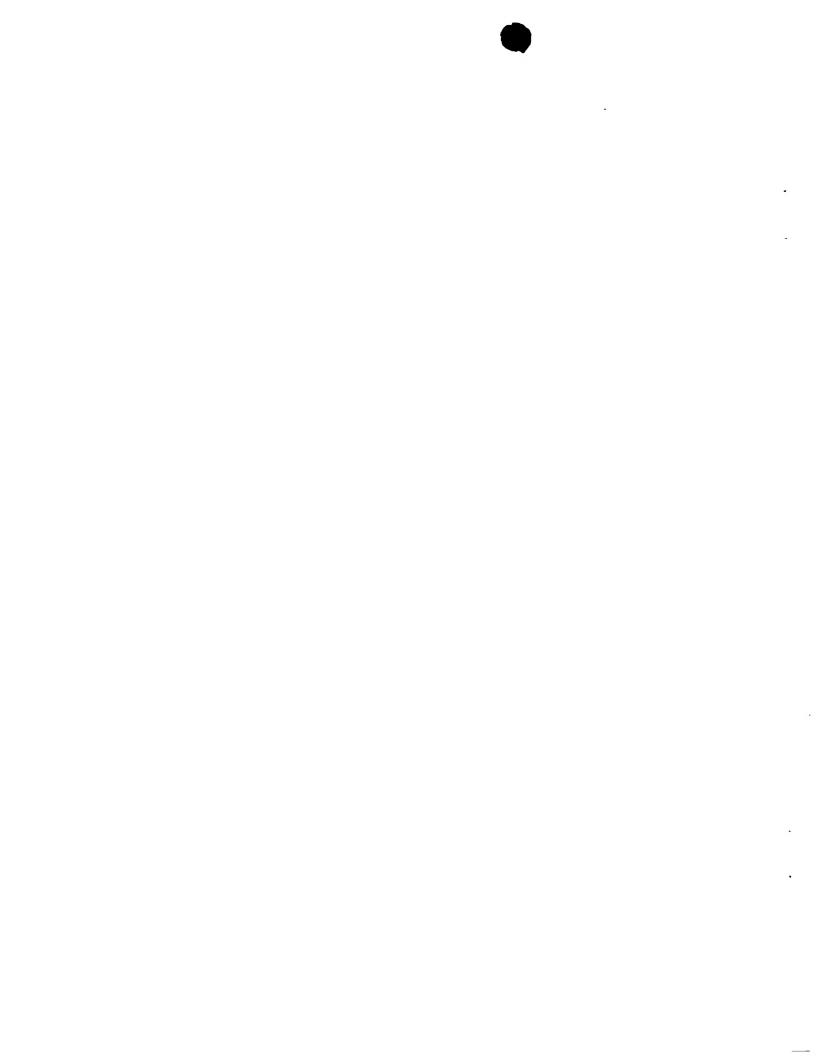
14. The method according to claim 12 or 13, wherein the processing tool is an optical receiver.

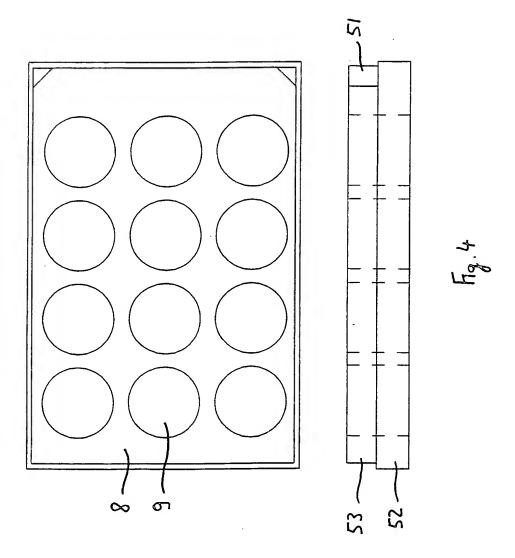
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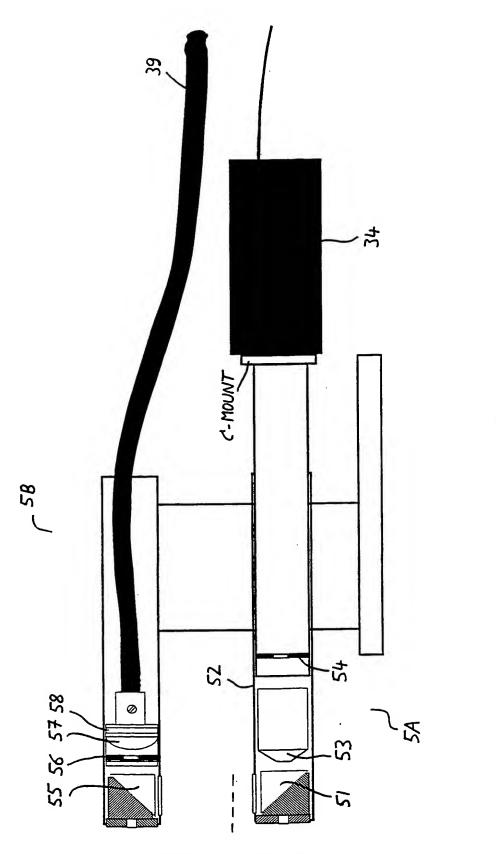
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ional Application No PCT/EP 00/06245

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 G01N35/04 G02B21/00

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)  $IPC \ 7 \quad B65D \quad G02B \quad G01N$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## EPO-Internal

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 98 52047 A (AURORA BIOSCIENCES CORP) 19 November 1998 (1998-11-19) page 84, paragraph 3 -page 85, paragraph 2; figure 15	1,2
A	WO 98 59033 A (SANKYO CO ;ICHIKAWA MASATO (JP); INABA KAZUHIRO (JP); NITTETSU MIN) 30 December 1998 (1998-12-30) figure 6	2
Α	DE 298 06 303 U (ARCHYTAS AUTOMATION GMBH) 3 September 1998 (1998-09-03) page 11, last paragraph -page 12, paragraph 2; figure 1 page 16, line 10 - line 32; figure 4 -/	1,2,6,10

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
*Special categories of cited documents:  "A" document defining the general state of the art which is not considered to be of particular relevance  "E" earlier document but published on or after the international filing date  "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  "O" document referring to an oral disclosure, use, exhibition or other means  "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  "&" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the International search report
25 October 2000	02/11/2000
Name and mailing address of the ISA	Authorized officer
European Patent Office, P.B. 5818 Patentiaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo ni, Fax: (+31–70) 340–3016	Hocquet, A

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## INTERNATIONAL SEARCH REPORT

Inte Ional Application No PCT/EP 00/06245

		PCI/EP OC	J/ U0245
	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
A	EP 0 245 920 A (CAMBRIDGE LIFE SCIENCES) 19 November 1987 (1987-11-19) column 4, line 40 -column 6, line 13; figures 2,3		6,7
Α	US 5 386 318 A (KUEHNERT JUERGEN ET AL) 31 January 1995 (1995-01-31) cited in the application column 3, line 48 -column 4, line 42; figures		1,2,7,8, 10
A	WO 97 39348 A (ALPHA SCIENT CORP) 23 October 1997 (1997-10-23) cited in the application page 7, last paragraph -page 8, paragraph 3; figures 3-6		1,2,10
A	DE 297 12 535 U (BODENSEEWERK PERKIN ELMER CO) 18 September 1997 (1997-09-18) cited in the application page 6, paragraph 2; figure 1 page 7, paragraph 3; figures 2,4		1,2,10
A	WO 93 12430 A (BAXTER DIAGNOSTICS INC) 24 June 1993 (1993-06-24) page 37, line 23 -page 39, line 33; figures 26-31		1,2,10
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information on patent family members

International Application No PC1/EP 00/06245

	ent document n search report		Publication date		atent family member(s)	Publication date
WO 9	9852047	A	19-11-1998	US AU EP	5985214 A 7478398 A 1010009 A	16-11-1999 08-12-1998 21-06-2000
WO 9	9859033	A	30-12-1998	JP AU EP	11009260 A 8037798 A 1018544 A	19-01-1999 04-01-1999 12-07-2000
DE 2	29806303	U	03-09-1998	NONE	<del></del>	
EP C	0245920	A	19–11–1987	GB AU DK FI HU JP US	2190195 A 6985487 A 236487 A 872019 A 43733 A,8 63061142 A 4810096 A	11-11-1987 12-11-1987 10-11-1987 10-11-1987 30-11-1987 17-03-1988 07-03-1989
US 5	5386318	Α	31-01-1995	DE WO	4131360 A 9306516 A	25-03-1993 01-04-1993
WO 9	9739348	A	23-10-1997	AU	2600897 A	07-11-1997
DE 2	29712535	U	18-09-1997	NONE		
WO 9	9312430	A	24-06-1993	AU CA EP JP	3333493 A 2101951 A 0572640 A 6507976 T	19-07-1993 19-07-1992 08-12-1993 08-09-1994

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